

FILE 'USPAT' ENTERED AT 13:47:52 ON 31 JUL 1998

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* * * * *
*   W E L C O M E   T O   T H E
*   U . S .   P A T E N T   T E X T   F I L E
* * * * *
```

=> s collagen(la)(matrix or gel# or implant#)

11664 COLLAGEN  
148931 MATRIX  
163494 GEL#  
23366 IMPLANT#

L1 988 COLLAGEN(1A)(MATRIX OR GEL# OR IMPLANT#)

=> s (BONE(W)MORPHOGEN?) or BMP? or (OSTEOGENIC(W)(PROTEIN? OR POLYPEPTIDE?))

31219 BONE  
876 MORPHOGEN?  
431 BONE(W)MORPHOGEN?  
672 BMP?  
659 OSTEOGENIC  
78334 PROTEIN?  
17867 POLYPEPTIDE?  
93 OSTEOGENIC(W)(PROTEIN? OR POLYPEPTIDE?)

L2 834 (BONE(W)MORPHOGEN?) OR BMP? OR (OSTEOGENIC(W)(PROTEIN? OR P  
OLY

PEPTIDE?))

=> s TGFbeta## or (TGF(W)BETA##) or ((TRANSFORMING(W)GROWTH(W)FACTOR#)(1A)beta##)

3 TGFbeta##  
1598 TGF  
170609 BETA##  
1030 TGF(W)BETA##  
27776 TRANSFORMING  
137791 GROWTH  
415655 FACTOR#  
170609 BETA##

L3 741 (TRANSFORMING(W)GROWTH(W)FACTOR#)(1A)BETA##  
1255 TGFbeta## OR (TGF(W)BETA##) OR ((TRANSFORMING(W)GROWTH(W)FA

CTO R#)(1A)BETA##)

=> s l1 and l2

L4 101 L1 AND L2

=> s l1(p)l2

L5 21 L1(P)L2

=> s binding or binder

101283 BINDING  
92565 BINDER  
L6 178310 BINDING OR BINDER

=> s l5(p)l6

L7 0 L5(P)L6

=> s viscous or viscosity

84386 VISCIOUS  
175038 VISCOSITY  
L8 217752 VISCIOUS OR VISCOSITY

=> s l5 and l8

L9 9 L5 AND L8

=> s ?cellulose

<-----User Break----->

L10 144531 ?CELLULOSE

=>

u

=>

=>

YOU HAVE RECEIVED 3 CONSECUTIVE ARROW PROMPTS (=>)  
The arrow (=>) is the system prompt, where you enter a command.  
If you need an explanation of system commands, files, formats, etc.,  
enter HELP and the name of the item you want explained at an arrow  
prompt (=>). The '?' can be used as a synonym for HELP at any  
prompt. Enter HELP COMMANDS for a list of commands that can be used  
in this file. Enter HELP MESSAGES for a list of online explanations  
that are available.

Help is available at any prompt, and after any error message.  
Enter HELP or '?' at a prompt to see an explanation of the options.

After an error message, enter HELP or '?' at the next prompt and you will receive a more detailed explanation of the error and how to correct it.

Automatic help is also available. To turn off the automatic help feature, enter SET AUHELP NONE at an arrow prompt. When the SET parameter AUHELP is ON, you will automatically receive help following an error message. To receive automatic help after 2 consecutive error messages or 3 consecutive prompts, enter SET AUTOHELP OFF. For more information on the SET parameter AUHELP, enter HELP SET AUHELP at the arrow prompt (=>).

For a list of commands, enter HELP COMMANDS. For a list of online explanations, enter HELP MESSAGES.

IF YOU REQUIRE FURTHER HELP, PLEASE CONTACT YOUR LOCAL HELP DESK

=> d his

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(FILE 'USPAT' ENTERED AT 13:47:52 ON 31 JUL 1998)
L1      988 S COLLAGEN(1A) (MATRIX OR GEL# OR IMPLANT#)
L2      834 S (BONE(W)MORPHOGEN?) OR BMP? OR (OSTEOGENIC(W) (PROTEIN? O
R P
L3      1255 S TGF BETA## OR (TGF(W)BETA##) OR ((TRANSFORMING(W)GROWTH(W
)FA
L4      101 S L1 AND L2
L5      21 S L1(P)L2
L6      178310 S BINDING OR BINDER
L7      0 S L5(P)L6
L8      217752 S VISCOUS OR VISCOSITY
L9      9 S L5 AND L8
L10     144531 S ?CELLULOSE
```

=> s 15(p)?cellulose

```
144531 ?CELLULOSE
L11     0 L5(P)?CELLULOSE
```

=> s 15(p) (?cellulose or cellulosic)

```
144531 ?CELLULOSE
21781 CELLULOSIC
L12     0 L5(P) (?CELLULOSE OR CELLULOSIC)
```

=> s 15(2p) (?cellulose or cellulosic)

```
144531 ?CELLULOSE
21781 CELLULOSIC
L13     1 L5(2P) (?CELLULOSE OR CELLULOSIC)
```

=> d 1-

1. 5,645,591, Jul. 8, 1997, Synthetic bone matrix; Thangavel Kuberasampath, et al., 623/16, 66 [IMAGE AVAILABLE]

=> select ]

ENTER ANSWER SET L#, TERMSET L# or (L13):end

=> select

ENTER ANSWER SET L#, TERMSET L# or (L13):l13

ENTER ANSWER NUMBER OR RANGE (1-):1

ENTER DISPLAY FORMAT (TI) OR ?:pn

E1 THROUGH E1 ASSIGNED

=> s e1

```
L14     1 "5,645,591"/PN
        (5645591/PN)
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=> d his

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(FILE 'USPAT' ENTERED AT 13:47:52 ON 31 JUL 1998)
L1      988 S COLLAGEN(1A) (MATRIX OR GEL# OR IMPLANT#)
L2      834 S (BONE(W)MORPHOGEN?) OR BMP? OR (OSTEOGENIC(W) (PROTEIN? O
R P
L3      1255 S TGF BETA## OR (TGF(W)BETA##) OR ((TRANSFORMING(W)GROWTH(W
)FA
L4      101 S L1 AND L2
L5      21 S L1(P)L2
L6      178310 S BINDING OR BINDER
L7      0 S L5(P)L6
L8      217752 S VISCOUS OR VISCOSITY
L9      9 S L5 AND L8
L10     144531 S ?CELLULOSE
L11     0 S L5(P)?CELLULOSE
L12     0 S L5(P) (?CELLULOSE OR CELLULOSIC)
```

L13 1 S L5(2P) (?CELLULOSE OR CELLULOSIC)  
 SELECT  
 L13 1 PN  
 L14 1 S E1  
 => s l14 and (collagen(2p) (?CELLULOSE OR CELLULOSIC))  
 11664 COLLAGEN  
 144531 ?CELLULOSE  
 21781 CELLULOSIC  
 2529 COLLAGEN(2P) (?CELLULOSE OR CELLULOSIC)  
 L15 1 L14 AND (COLLAGEN(2P) (?CELLULOSE OR CELLULOSIC))  
 => d kwic  
 US PAT NO: \*\*5,645,591\*\* [IMAGE AVAILABLE] L15: 1 of 1  
 SUMMARY:  
 BSUM(15)  
 The \*\*collagen\*\*-GAG polymer is cross-linked to control the solubility and mechanical properties of the matrix. It has been determined that cross-linking the. . .  
 SUMMARY:  
 BSUM(16)  
 The invention is embodied as a method of growing bone by conduction including contacting a viable mammalian bone with the cross-linked \*\*collagen\*\*-GAG matrix. Bone conduction is the growth of bone from existing viable bone, and involves the migration of osteoblasts from the. . . to solidify the matrix when implanted in a mammal or when placed at 37.degree. C. A useful glue is methyl \*\*cellulose\*\*. The matrix solidifies substantially in the shape of the implanted matrix.  
 SUMMARY:  
 BSUM(18)  
 Another . . . of producing the osteogenic device which contains osteogenic protein. The method includes providing a porous matrix comprising a polymer of \*\*collagen\*\* and GAG cross-linked to an M.sub.c value of about 800 to about 60,000; and dispersing within the matrix an osteogenic. . .  
 DETDESC:  
 DETD(28)  
 Alternatively, . . . cobalt, or polymers such as polyglycolic acid or polylactic acid. Upon the addition of a heat-activated glue such as methyl \*\*cellulose\*\*, the material becomes solidified after implantation or when placed at 37.degree. C.  
 DETDESC:  
 DETD(29)  
 Thus, . . . and cell division. Hence, osteoblasts may be induced to migrate from viable bone to the material. In addition, the cross-linked \*\*collagen\*\*-GAG material has a negative surface charge which enhances cell attachment. Furthermore, osteoblasts synthesize fibronectin, a cellular adherence protein that binds \*\*collagen\*\*, thereby enhancing the ability of the migrating osteoblasts to adhere to the implant.  
 DETDESC:  
 DETD(31)  
 The . . . also comprise a molded, porous solid, or simply an aggregation of close-packed particles held in place by surrounding tissue. Insoluble \*\*collagen\*\* or inert polymers added to the \*\*collagen\*\*-GAG-osteogenic protein particles may increase the density of the device. In addition, a glue or solidifying agent including methyl \*\*cellulose\*\*, (e.g., Methocel, Dow Chemical Co.), may be added. It is preferable to shape the matrix into the desired form of. . .  
 DETDESC:  
 DETD(32)  
 The . . . and evaluated histologically for evidence of bone formation. FIG. 1 demonstrates that only mesenchymal cells will be associated with a \*\*collagen\*\*-GAG implant that does not include osteogenic protein, while FIG. 2 shows the ultimate development of endochondral bone in an implant. . .  
 CLAIMS:  
 CLMS(10)

10. The method of claim 9 wherein said glue comprises methyl  
\*\*cellulose\*\*.

CLAIMS:

CLMS(11)

11. . . . method of growing mammalian bone by conduction comprising  
the steps of:

(A) providing a porous matrix comprising a polymer of \*\*collagen\*\* and  
glycosaminoglycan cross-linked to an Mc value of about 800 to about  
60,000 and a glue comprising methyl \*\*cellulose\*\* in an amount  
sufficient to solidify said matrix when implanted in a mammal; and  
(b) contacting viable mammalian bone at the. . .

=> d bib date ab

US PAT NO: \*\*5,645,591\*\* [IMAGE AVAILABLE] L15: 1 of 1  
DATE ISSUED: Jul. 8, 1997  
TITLE: Synthetic bone matrix  
INVENTOR: Thangavel Kuberasampath, Medway, MA  
Lawrence Berlowitz Tarrant, Harvard, MA  
ASSIGNEE: Stryker Corporation, Natick, MA (U.S. corp.)  
APPL-NO: 07/529,852  
DATE FILED: May 29, 1990  
ART-UNIT: 338  
PRIM-EXMR: David Isabella  
LEGAL-REP: Testa, Hurwitz & Thibault

L15: 1 of 1

TITLE: Synthetic bone matrix  
US PAT NO: \*\*5,645,591\*\* DATE ISSUED: Jul. 8, 1997  
[IMAGE AVAILABLE]  
APPL-NO: 07/529,852 DATE FILED: May 29, 1990

ABSTRACT:

Disclosed is an osteogenic device capable of inducing the formation of  
endochondral bone in a shape conforming substantially to the shape of the  
device when implanted in a mammalian host. The device includes an  
osteogenic protein dispersed within a porous matrix comprising a polymer  
of collagen and glycosaminoglycan cross-linked to an M.sub.c value of  
about 800 to about 60,000. Also disclosed are a method of inducing  
mammalian bone growth, and a method of inducing conductive bone growth  
from viable mammalian bone.

=> d his

(FILE 'USPAT' ENTERED AT 13:47:52 ON 31 JUL 1998)

L1 988 S COLLAGEN(1A) (MATRIX OR GEL# OR IMPLANT#)  
L2 834 S (BONE(W)MORPHOGEN?) OR BMP? OR (OSTEOGENIC(W) (PROTEIN? O  
R P  
L3 1255 S TGFbeta## OR (TGF(W)BETA##) OR ((TRANSFORMING(W)GROWTH(W  
)FA  
L4 101 S L1 AND L2  
L5 21 S L1(P)L2  
L6 178310 S BINDING OR BINDER  
L7 0 S L5(P)L6  
L8 217752 S VISCOUS OR VISCOSITY  
L9 9 S L5 AND L8  
L10 144531 S ?CELLULOSE  
L11 0 S L5(P)?CELLULOSE  
L12 0 S L5(P) (?CELLULOSE OR CELLULOSIC)  
L13 1 S L5(2P) (?CELLULOSE OR CELLULOSIC)  
SELECT  
L13 1 PN  
L14 1 S E1  
L15 1 S L14 AND (COLLAGEN(2P) (?CELLULOSE OR CELLULOSIC))

=> d bib date ab 1- 19

US PAT NO: 5,550,188 [IMAGE AVAILABLE] L9: 1 of 9  
DATE ISSUED: Aug. 27, 1996  
TITLE: Polymer conjugates ophthalmic devices comprising  
collagen-polymer conjugates  
INVENTOR: Woonza Rhee, Palo Alto, CA  
Donald G. Wallace, Menlo Park, CA  
Alan S. Michaels, Boston, MA  
Ramon A. Burns, Jr., Fremont, CA  
Louis Fries, Los Altos, CA  
Frank DeLustro, Belmont, CA  
Hanne Bentz, Newark, CA  
ASSIGNEE: Collagen Corporation, Palo Alto, CA (U.S. corp.)  
APPL-NO: 08/478,510  
DATE FILED: Jun. 7, 1995  
ART-UNIT: 127  
PRIM-EXMR: Nathan M. Nutter  
LEGAL-REP: Morrison & Foerster

TITLE: Polymer conjugates ophthalmic devices comprising collagen-polymer conjugates  
 US PAT NO: 5,550,188 DATE ISSUED: Aug. 27, 1996  
 [IMAGE AVAILABLE]  
 APPL-NO: 08/478,510 DATE FILED: Jun. 7, 1995  
 REL-US-DATA: Division of Ser. No. 368,874, Jan. 5, 1995, Pat. No. 5,446,051, which is a division of Ser. No. 198,128, Feb. 17, 1994, Pat. No. 5,413,791, which is a division of Ser. No. 922,541, Jul. 30, 1992, Pat. No. 5,328,955, which is a continuation-in-part of Ser. No. 433,441, Nov. 14, 1989, Pat. No. 5,162,430, which is a continuation-in-part of Ser. No. 274,071, Nov. 21, 1988, abandoned.

ABSTRACT:  
 Pharmaceutically acceptable, non-immunogenic compositions are formed by covalently binding atelopeptide collagen to pharmaceutically pure, synthetic, hydrophilic polymers via specific types of chemical bonds to provide collagen/polymer conjugates. The atelopeptide collagen can be type I, type II or type III and may be fibrillar or non-fibrillar. The synthetic hydrophilic polymer may be polyethylene glycol and derivatives thereof having a weight average molecular weight over a range of from about 100 to about 20,000. The compositions may include other components such as liquid, pharmaceutically acceptable, carriers to form injectable formulations, and/or biologically active proteins such as growth factors. The collagen-polymer conjugates of the invention generally contain large amounts of water when formed. The conjugates can be dehydrated to form a relatively solid object. The dehydrated, solid object can be ground into particles which can be suspended in a non-aqueous fluid such as an oil and injected into a living being for the purpose of providing soft tissue augmentation. Once in place, the particles rehydrate and expand in size five fold or more.

US PAT NO: 5,475,052 [IMAGE AVAILABLE]  
 DATE ISSUED: Dec. 12, 1995  
 TITLE: Collagen-synthetic polymer matrices prepared using a multiple step reaction  
 INVENTOR: Woonza M. Rhee, Palo Alto, CA  
 Richard A. Berg, Los Altos, CA  
 ASSIGNEE: Collagen Corporation, Palo Alto, CA (U.S. corp.)  
 APPL-NO: 08/236,769  
 DATE FILED: May 2, 1994  
 ART-UNIT: 127  
 PRIM-EXMR: Nathan M. Nutter  
 LEGAL-REP: Kathi Rafayko

TITLE: Collagen-synthetic polymer matrices prepared using a multiple step reaction  
 US PAT NO: 5,475,052 DATE ISSUED: Dec. 12, 1995  
 [IMAGE AVAILABLE]  
 APPL-NO: 08/236,769 DATE FILED: May 2, 1994  
 REL-US-DATA: Continuation-in-part of Ser. No. 198,128, Feb. 17, 1994, which is a division of Ser. No. 922,541, Jul. 30, 1992, Pat. No. 5,328,955, which is a continuation-in-part of Ser. No. 433,441, Nov. 14, 1989, Pat. No. 5,162,430, Nov. 10, 1992, which is a continuation-in-part of Ser. No. 274,071, Nov. 21, 1988, abandoned.

ABSTRACT:  
 The present invention discloses collagen-synthetic polymer matrices which are prepared using a multiple step reaction. The first step of the reaction generally involves reacting collagen with a functionally activated synthetic hydrophilic polymer to form a collagen-synthetic polymer matrix. The synthetic hydrophilic polymer may be mono- or multifunctionally activated, but is preferably difunctionally activated, resulting in the formation of a crosslinked collagen matrix. The second step comprises modifying the collagen-synthetic polymer matrix according to one or more of the following methods: further crosslinking the matrix using a multifunctionally activated synthetic polymer, conjugating the matrix using a monofunctionally activated synthetic polymer, coupling biologically active molecules or glycosaminoglycans to the matrix, crosslinking the matrix using conventional chemical crosslinking agents, or modifying the collagen in the matrix by means of various chemical reactions. An optional third step may include further modification of the collagen-synthetic polymer matrix by covalently binding, for example, biologically active molecules or glycosaminoglycans to the matrix by means of available active groups on the synthetic hydrophilic polymers. Collagen-synthetic polymer matrices prepared according to the methods of the present invention have very low immunogenicity and can therefore be used to prepare biocompatible implants for use in a variety of medical applications.

US PAT NO: 5,446,091 [IMAGE AVAILABLE]  
 DATE ISSUED: Aug. 29, 1995  
 TITLE: Collagen-polymer conjugates containing an ether linkage  
 INVENTOR: Woonza Rhee, Palo Alto, CA  
 Donald G. Wallace, Menlo Park, CA  
 Alan S. Michaels, Boston, MA  
 Ramon A. Burns, Jr., Fremont, CA

Louis Fries, Los Altos, CA  
Frank DeLustro, Belmont, CA  
Hanne Bentz, Newark, CA  
ASSIGNEE: Collagen Corporation, Palo Alto, CA (U.S. corp.)  
APPL-NO: 08/368,874  
DATE FILED: Jan. 5, 1995  
ART-UNIT: 153  
PRIM-EXMR: Nathan M. Nutter  
LEGAL-REP: Morrison & Foerster

L9: 3 of 9

TITLE: Collagen-polymer conjugates containing an ether linkage  
US PAT NO: 5,446,091 DATE ISSUED: Aug. 29, 1995  
[IMAGE AVAILABLE]  
APPL-NO: 08/368,874 DATE FILED: Jan. 5, 1995  
REL-US-DATA: Division of Ser. No. 198,128, Feb. 17, 1994, Pat. No. 5,413,791, which is a division of Ser. No. 922,541, Jun. 30, 1992, Pat. No. 5,328,955, Jul. 12, 1994, which is a continuation-in-part of Ser. No. 433,441, Nov. 14, 1989, Pat. No. 5,162,430, Nov. 10, 1992, which is a continuation-in-part of Ser. No. 274,071, Nov. 21, 1988, abandoned.

ABSTRACT:

Pharmaceutically acceptable, non-immunogenic compositions are formed by covalently binding atelopeptide collagen to pharmaceutically pure, synthetic, hydrophilic polymers via specific types of chemical bonds to provide collagen/polymer conjugates. The atelopeptide collagen can be type I, type II or type III and may be fibrillar or non-fibrillar. The synthetic hydrophilic polymer may be polyethylene glycol and derivatives thereof having a weight average molecular weight over a range of from about 100 to about 20,000. The compositions may include other components such as liquid, pharmaceutically acceptable, carriers to form injectable formulations, and/or biologically active proteins such as growth factors. The collagen-polymer conjugates of the invention generally contain large amounts of water when formed. The conjugates can be dehydrated to form a relatively solid object. The dehydrated, solid object can be ground into particles which can be suspended in a non-aqueous fluid such as an oil and injected into a living being for the purpose of providing soft tissue augmentation. Once in place, the particles rehydrate and expand in size five fold or more.

US PAT NO: 5,413,989 [IMAGE AVAILABLE] L9: 4 of 9  
DATE ISSUED: May 9, 1995  
TITLE: Method and activin compositions for inducing bone growth  
INVENTOR: Yasushi Ogawa, Pacifica, CA  
David K. Schmidt, Santa Cruz, CA  
Rosa Armstrong, Palo Alto, CA  
Ranga Nathan, Newark, CA  
Andrea Y. Thompson, Mountain View, CA  
Saeid M. Seyedin, Saratoga, CA  
ASSIGNEE: Celtrix Pharmaceuticals, Inc., Santa Clara, CA (U.S. corp.)  
APPL-NO: 08/056,469  
DATE FILED: May 3, 1993  
ART-UNIT: 181  
PRIM-EXMR: Jill A. Warden  
ASST-EXMR: Carol A. Salata  
LEGAL-REP: Morrison & Foerster

L9: 4 of 9

TITLE: Method and activin compositions for inducing bone growth  
US PAT NO: 5,413,989 DATE ISSUED: May 9, 1995  
[IMAGE AVAILABLE] DISCL-DATE: May 4, 2010  
APPL-NO: 08/056,469 DATE FILED: May 3, 1993  
REL-US-DATA: Continuation of Ser. No. 655,313, Feb. 14, 1991, Pat. No. 5,208,219.

ABSTRACT:

Activin is administered systemically and locally to induce the growth of mature bone. Activin enhances the level of bone formation and the quality of the bone formed when administered locally with BMP or bone marrow. Administration of activin by subcutaneous route promotes systemic increase in the bone mass.

US PAT NO: 5,413,791 [IMAGE AVAILABLE] L9: 5 of 9  
DATE ISSUED: May 9, 1995  
TITLE: Collagen-polymer conjugates  
INVENTOR: Woonza Rhee, Palo Alto, CA  
Donald G. Wallace, Menlo Park, CA  
Alan S. Michaels, Boston, MA  
Ramon A. Burns, Jr., Fremont, CA  
Louis Fries, Los Altos, CA  
Frank DeLustro, Belmont, CA  
Hanne Bentz, Newark, CA  
ASSIGNEE: Collagen Corporation, Palo Alto, CA (U.S. corp.)  
APPL-NO: 08/198,128  
DATE FILED: Feb. 17, 1994  
ART-UNIT: 153  
PRIM-EXMR: Nathan M. Nutter  
LEGAL-REP: Morrison & Foerster

TITLE: Collagen-polymer conjugates  
 US PAT NO: 5,413,791 DATE ISSUED: May 9, 1995  
 [IMAGE AVAILABLE]  
 APPL-NO: 08/198,128 DATE FILED: Feb. 17, 1994  
 REL-US-DATA: Division of Ser. No. 922,541, Jul. 30, 1992, Pat. No. 5,328,955, which is a continuation-in-part of Ser. No. 433,441, Nov. 14, 1989, Pat. No. 5,162,430, which is a continuation-in-part of Ser. No. 274,071, Nov. 21, 1988, abandoned.

ABSTRACT:  
 Pharmaceutically acceptable, non-immunogenic compositions are formed by covalently binding atelopeptide collagen to pharmaceutically pure, synthetic, hydrophilic polymers via specific types of chemical bonds to provide collagen/polymer conjugates. The atelopeptide collagen can be type I, type II or type III and may be fibrillar or non-fibrillar. The synthetic hydrophilic polymer may be polyethylene glycol and derivatives thereof having a weight average molecular weight over a range of from about 100 to about 20,000. The compositions may include other components such as liquid, pharmaceutically acceptable, carriers to form injectable formulations, and/or biologically active proteins such as growth factors. The collagen-polymer conjugates of the invention generally contain large amounts of water when formed. The conjugates can be dehydrated to form a relatively solid object. The dehydrated, solid object can be ground into particles which can be suspended in a non-aqueous fluid such as an oil and injected into a living being for the purpose of providing soft tissue augmentation. Once in place, the particles rehydrate and expand in size five fold or more.

US PAT NO: 5,328,955 [IMAGE AVAILABLE]  
 DATE ISSUED: Jul. 12, 1994  
 TITLE: Collagen-polymer conjugates  
 INVENTOR: Woonza Rhee, Palo Alto, CA  
 Donald G. Wallace, Menlo Park, CA  
 Alan S. Michaels, Boston, MA  
 Ramon A. Burns, Jr., Fremont, CA  
 Louis Fries, Los Altos, CA  
 Frank DeLustro, Belmont, CA  
 Hanne Bentz, Newark, CA  
 ASSIGNEE: Collagen Corporation, Palo Alto, CA (U.S. corp.)  
 APPL-NO: 07/922,541  
 DATE FILED: Jul. 30, 1992  
 ART-UNIT: 153  
 PRIM-EXMR: Nathan M. Nutter  
 LEGAL-REP: Karl Bozicevic

TITLE: Collagen-polymer conjugates  
 US PAT NO: 5,328,955 DATE ISSUED: Jul. 12, 1994  
 [IMAGE AVAILABLE]  
 APPL-NO: 07/922,541 DATE FILED: Jul. 30, 1992  
 REL-US-DATA: Continuation-in-part of Ser. No. 433,441, Nov. 14, 1989, Pat. No. 5,162,430, which is a continuation-in-part of Ser. No. 274,071, Nov. 21, 1988, abandoned.

ABSTRACT:  
 Pharmaceutically acceptable, non-immunogenic compositions are formed by covalently binding atelopeptide collagen to pharmaceutically pure, synthetic, hydrophilic polymers via specific types of chemical bonds to provide collagen/polymer conjugates. The atelopeptide collagen can be type I, type II or type III and may be fibrillar or non-fibrillar. The synthetic hydrophilic polymer may be polyethylene glycol and derivatives thereof having a weight average molecular weight over a range of from about 100 to about 20,000. The compositions may include other components such as liquid, pharmaceutically acceptable, carriers to form injectable formulations, and/or biologically active proteins such as growth factors. The collagen-polymer conjugates of the invention generally contain large amounts of water when formed. The conjugates can be dehydrated to form a relatively solid object. The dehydrated, solid object can be ground into particles which can be suspended in a non-aqueous fluid such as an oil and injected into a living being for the purpose of providing soft tissue augmentation. Once in place, the particles rehydrate and expand in size five fold or more.

US PAT NO: 5,308,889 [IMAGE AVAILABLE]  
 DATE ISSUED: May 3, 1994  
 TITLE: Dehydrated collagen-polymer strings  
 INVENTOR: Woonza Rhee, Palo Alto, CA  
 Louis Fries, Los Altos, CA  
 Ramesh Damani, Mountain View, CA  
 Kimberly McCullough, Hayward, CA  
 Frank DeLustro, Belmont, CA  
 ASSIGNEE: Collagen Corporation, Palo Alto, CA (U.S. corp.)  
 APPL-NO: 07/984,197  
 DATE FILED: Dec. 2, 1992  
 ART-UNIT: 153  
 PRIM-EXMR: Nathan M. Nutter  
 LEGAL-REP: Karl Bozicevic

TITLE: Dehydrated collagen-polymer strings  
 US PAT NO: 5,308,889 DATE ISSUED: May 3, 1994  
 [IMAGE AVAILABLE]  
 APPL-NO: 07/984,197 DATE FILED: Dec. 2, 1992  
 REL-US-DATA: Continuation-in-part of Ser. No. 922,541, Jul. 30, 1992,  
 which is a continuation-in-part of Ser. No. 433,441,  
 Nov. 14, 1989, Pat. No. 5,162,430, Nov. 10, 1992, which  
 is a continuation-in-part of Ser. No. 274,071, Nov. 21,  
 1988, abandoned.

## ABSTRACT:

Medical articles in the form of strings are formed by covalently binding collagen to pharmaceutically pure, synthetic, hydrophilic polymers via specific types of chemical bonds to provide collagen/polymer conjugate formulations which are extruded to make the strings. The collagen may be recombinantly produced human collagen or collagen extracted from any source, such as a bovine source or human placenta, and purified and can be of various types and may be fibrillar or non-fibrillar. The synthetic hydrophilic polymer may be polyethylene glycol and derivatives thereof having an average molecular weight over a range of from about 100 to about 20,000. The string can be designed to incorporate other components such as fluid, pharmaceutically acceptable carriers to form injectable formulations, and/or biologically active proteins such as growth factors or cytokines. The strings contain large amounts of water when extruded and may then be dehydrated to form relatively solid but flexible strings. The strings can be injected into a living being for the purpose of providing soft tissue augmentation. Once in place, the strings rehydrate and expand in size five fold or more. Aqueous solution can be provided to enhance the rate of rehydration. The strings can also be used to suture wounds which strings can be chemically designed to dissolve in situ.

US PAT NO: 5,292,802 [IMAGE AVAILABLE]  
 DATE ISSUED: Mar. 8, 1994  
 TITLE: Collagen-polymer tubes for use in vascular surgery  
 INVENTOR: Woonza Rhee, Palo Alto, CA  
 Kimberly McCullough, Hayward, CA  
 ASSIGNEE: Collagen Corporation, Palo Alto, CA (U.S. corp.)  
 APPL-NO: 07/985,680  
 DATE FILED: Dec. 2, 1992  
 ART-UNIT: 153  
 PRIM-EXMR: Nathan M. Nutter  
 LEGAL-REP: Karl Bozicevic

TITLE: Collagen-polymer tubes for use in vascular surgery  
 US PAT NO: 5,292,802 DATE ISSUED: Mar. 8, 1994  
 [IMAGE AVAILABLE]  
 APPL-NO: 07/985,680 DATE FILED: Dec. 2, 1992  
 REL-US-DATA: Continuation-in-part of Ser. No. 922,541, Jul. 30, 1992,  
 which is a continuation-in-part of Ser. No. 433,441,  
 Nov. 14, 1989, Pat. No. 5,162,430, Nov. 10, 1992, which  
 is a continuation-in-part of Ser. No. 274,071, Nov. 21,  
 1988, abandoned.

## ABSTRACT:

Medical articles in the form of tubes are formed by covalently binding collagen to pharmaceutically pure, synthetic, hydrophilic polymers via specific types of chemical bonds to provide collagen/polymer conjugate formulations which are used to make the tubes. The collagen may be recombinantly produced human collagen or collagen extracted from any source, such as a bovine or human placental source, and purified and can be type I, type II or type III and may be fibrillar or non-fibrillar. The synthetic hydrophilic polymer may be polyethylene glycol and derivatives thereof having a weight average molecular weight over a range of from about 100 to about 20,000. The tube can be designed to incorporate other components such as liquid, pharmaceutically acceptable, carriers, and/or biologically active proteins such as growth factors or cytokines. The tubes contain large amounts of water when extruded and then may be dehydrated to form relatively solid but flexible tubes which can be easily stored. The tubes can be surgically implanted and attached to, or implanted within, a channel in a mammal for the purpose of repairing the channel. The tubes can be used to repair a wide range of different types of channels including but not limited to veins and arteries.

US PAT NO: 5,208,219 [IMAGE AVAILABLE]  
 DATE ISSUED: May 4, 1993  
 TITLE: Method for inducing bone growth  
 INVENTOR: Yasushi Ogawa, Pacifica, CA  
 David K. Schmidt, Santa Cruz, CA  
 Rosa Armstrong, Palo Alto, CA  
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 Andrea Y. Thompson, Mountain View, CA  
 Saeid M. Seyedin, Saratoga, CA  
 ASSIGNEE: Celtrix Pharmaceuticals Inc., Santa Clara, CA (U.S. corp.)  
 APPL-NO: 07/655,313  
 DATE FILED: Feb. 14, 1991  
 ART-UNIT: 181  
 PRIM-EXMR: F. T. Moezie  
 LEGAL-REP: Morrison & Foerster



TITLE: Method for inducing bone growth  
 US PAT NO: 5,208,219 DATE ISSUED: May 4, 1993  
 [IMAGE AVAILABLE]  
 APPL-NO: 07/655,313 DATE FILED: Feb. 14, 1991

## ABSTRACT:

Activin is administered systemically and locally to induce the growth of mature bone. Activin enhances the level of bone formation and the quality of the bone formed when administered locally with BMP or bone marrow. Administration of activin by subcutaneous route promotes systemic increase in the bone mass.

=> save all a08822186/1

'A08822186/L' IN USE

REPLACE OLD DEFINITION? Y/(N):n

=> append all a08822186/1

'APPEND' IS NOT A RECOGNIZED COMMAND

=> act a08822186/1

L16 ( 19488)SEA FILE=USPAT ((OSTEOGENIC OR (BONE MORPHOGENETIC)) (W)PRO  
 TEI  
 L17 ( 11949)SEA FILE=USPAT COLLAGEN?  
 L18 ( 188)SEA FILE=USPAT DEMINERALIZED BONE  
 L19 ( 4637)SEA FILE=USPAT ?APATITE OR ?APATITES  
 L20 ( 138488)SEA FILE=USPAT PHOSPHATE?  
 L21 ( 87)SEA FILE=USPAT L17 (P) L18  
 L22 ( 19)SEA FILE=USPAT L21 (P) L19  
 L23 ( 3)SEA FILE=USPAT L22(P)L20  
 L24 ( 2)SEA FILE=USPAT L23 AND L16  
 L25 ( 20323)SEA FILE=USPAT MANNITOL  
 L26 ( 12895)SEA FILE=USPAT DEXTRAN OR DEXTRANS  
 L27 ( 1214)SEA FILE=USPAT WHITE PETROLATUM  
 L28 ( 6784)SEA FILE=USPAT SESAME OIL  
 L29 ( 4052)SEA FILE=USPAT CELLULOSES  
 L30 ( 14013)SEA FILE=USPAT L26 OR L27  
 L31 ( 1214)SEA FILE=USPAT L27(P)L30  
 L32 ( 31221)SEA FILE=USPAT L25 OR L26  
 L33 ( 28)SEA FILE=USPAT L27(P)L32  
 L34 ( 1068)SEA FILE=USPAT L28(P)L32  
 L35 ( 481)SEA FILE=USPAT L29(P)L32  
 L36 ( 0)SEA FILE=USPAT L33(P)L28  
 L37 ( 0)SEA FILE=USPAT L33(P)L29  
 L38 ( 47)SEA FILE=USPAT L16 AND (L33 OR L34 OR L35)

=> d his

(FILE 'USPAT' ENTERED AT 13:47:52 ON 31 JUL 1998)  
 L1 988 S COLLAGEN(1A) (MATRIX OR GEL# OR IMPLANT#)  
 L2 834 S (BONE(W)MORPHOGEN?) OR BMP? OR (OSTEOGENIC(W) (PROTEIN? O  
 R P  
 L3 1255 S TGF BETA## OR (TGF(W)BETA##) OR ((TRANSFORMING(W)GROWTH(W  
 )FA  
 L4 101 S L1 AND L2  
 L5 21 S L1(P)L2  
 L6 178310 S BINDING OR BINDER  
 L7 0 S L5(P)L6  
 L8 217752 S VISCOUS OR VISCOSITY  
 L9 9 S L5 AND L8  
 L10 144531 S ?CELLULOSE  
 L11 0 S L5(P)?CELLULOSE  
 L12 0 S L5(P) (?CELLULOSE OR CELLULOSIC)  
 L13 1 S L5(2P) (?CELLULOSE OR CELLULOSIC)  
 SELECT  
 L13 1 PN  
 L14 1 S E1  
 L15 1 S L14 AND (COLLAGEN(2P) (?CELLULOSE OR CELLULOSIC))  
 ACT A08822186/L  
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L16 ( 19488)SEA FILE=USPAT ((OSTEOGENIC OR (BONE MORPHOGENETIC)) (W)PRO  
 TEI  
 L17 ( 11949)SEA FILE=USPAT COLLAGEN?  
 L18 ( 188)SEA FILE=USPAT DEMINERALIZED BONE  
 L19 ( 4637)SEA FILE=USPAT ?APATITE OR ?APATITES  
 L20 ( 138488)SEA FILE=USPAT PHOSPHATE?  
 L21 ( 87)SEA FILE=USPAT L17 (P) L18  
 L22 ( 19)SEA FILE=USPAT L21 (P) L19  
 L23 ( 3)SEA FILE=USPAT L22(P)L20  
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L33 ( 28)SEA FILE=USPAT L27(P)L32  
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L35 ( 481)SEA FILE=USPAT L29(P)L32  
L36 ( 0)SEA FILE=USPAT L33(P)L28  
L37 ( 0)SEA FILE=USPAT L33(P)L29  
L38 ( 47)SEA FILE=USPAT L16 AND (L33 OR L34 OR L35)  
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